subgluteal abscesses are recognized complications following a pudendal block,6 but their incidence, which is low, is unknown. It has been suggested that the incidence of hematoma formation and infection of the hematoma is kept low by compression of the paravaginal tissues by the fetal head.7(p351)

A gallium citrate Ga 67 scan may be effective in visualizing a retropsoas abscess.8 In this report, CT has been shown to be equally effective in visualizing such a retroperitoneal hematoma or abscess. The use of CT to diagnose retroperitoneal fluid collections was recently reviewed.9 Retroperitoneal hemorrhage at the psoas muscle has been found by CT imaging when it arises as a consequence of a bleeding diathesis or anticoagulation, a ruptured abdominal aortic aneurysm,9 or most frequently as a complication of femoral catheterization.10 This is the first report of the use of CT to diagnose retroperitoneal hemorrhage complicating pudendal block. It is the diagnostic method of choice in studying retroperitoneal collections. It also is the most cost-effective technique while providing the same information about this process as can be derived from magnetic resonance imaging.

Vascular injury in the course of pudendal block is not a rare occurrence (as evidenced by positive aspiration tests). Fortunately, most patients with this injury are asymptomatic and have no sequelae, for reasons cited earlier. Physicians must, however, always remain cognizant that retroperitoneal bleeding may be life-threatening because of the possible size of this space. A key to the presence of an infected retroperitoneal hematoma is a postpartum temperature elevation in a patient with a dropping hematocrit. On the other hand, postpartum fever, usually of a wildly spiking nature that does not respond to antibiotics alone and in a patient whose hematocrit is stable, may point to a septic pelvic thrombophlebitis. The latter disorder, which is also rare—1 to 5 per 10,000 vaginal deliveries11,12—may similarly be diagnosed by CT imaging of the pelvis. 13-15 Septic pelvic thrombophlebitis requires the addition of heparin for a therapeutic response. Differentiating these two disorders by CT imaging could prevent a catastrophic outcome because the treatment of a patient with heparin in whom there is an underlying vessel perforation is contraindicated. Therefore, if pudendal block is followed by an unexplained fever postpartum, with hip or abdominal pain, CT may help to detect an infected retroperitoneal hematoma or abscess. Either disorder may be life-threatening or severely debilitating4 and hence must be treated aggressively.

### REFERENCES

- 1. Gabbe S, Niebyl JR, Simpson JL: Obstetrics—Normal and Problem Pregnancies, 2nd edition. New York, NY, Churchill-Livingstone, 1991
- 2. Cunningham FG, MacDonald PC, Gant NF, Leveno KJ, Gilstrap LC III: Williams Obstetrics, 19th edition. Norwalk, Conn, Appleton & Lange, 1994
- 3. Bozynski ME, Rubarth LB, Patel JA: Lidocaine toxicity after maternal udendal anesthesia in a term infant with fetal distress. Am J Perinatol 1987; 4: 164-166
- 4. Svancarek W, Chirino O, Schaefer G Jr, Blythe JG: Retropsoas and subgluteal abscesses following paracervical and pudendal anesthesia. JAMA 1977; 237:892-894
- 5. Wenger DR, Gitchell RG: Severe infections following pudendal block anesthesia: Need for orthopaedic awareness. J Bone Joint Surg [Am] 1973; 55:202-207

- 6. Zador G. Lindmark G. Nilsson BA: Pudendal block in normal vaginal deliveries—Clinical efficacy, lidocaine concentrations in maternal and foetal blood, foetal and maternal acid-base values and influence on uterine activity. Acta Obstet Gynecol Scand (Suppl) 1974; 34:51-64
- 7. Bowes WA Jr: Clinical aspects of normal and abnormal labor, chap 35, In Creasy RK, Resnik R (Eds): Maternal Fetal Medicine—Principles and Practice, 3rd edition. Philadelphia, Pa, WB Saunders, 1994
- 8. Oster MW, Gelrud LG, Lotz MJ, et al: Psoas abscess localization by gallium scan in aplastic anemia. JAMA 1975; 232:377-379
- 9. Korobkin M, Silverman PM, Quint LE, Francis IR: CT of the extraperitoneal space: Normal anatomy and fluid collections. AJR Am J Roentgenol 1992; 159:933-941
- $10.\ Trerotola$  SO, Kuhlman JE, Fishman EK: Bleeding complications of femoral catheterization: CT evaluation. Radiology 1990; 174:37-40
- 11. Duff P, Gibbs RS: Pelvic vein thrombophlebitis: Diagnostic dilemma and therapeutic challenge. Obstet Gynecol Surv 1983; 38:365-37
- 12. Sweet RL, Ledger WJ: Puerperal infectious morbidity—A two year review. Am J Obstet Gynecol 1973; 117:1093-1100
- 13. Bahnson RR, Wendel EF, Vogelzang RL: Renal vein thrombosis following puerperal ovarian vein thrombophlebitis. Am J Obstet Gynecol 1985; 152:
- 14. Brown CEL, Lowe TW, Cunningham FG, Weinreb JC: Puerperal pelvic thrombophlebitis: Impact on diagnosis and treatment using x-ray computed tomography and magnetic resonance imaging. Obstet Gynecol 1986; 68:789-794
- 15. Angel JL, Knuppel RA: Computed tomography in diagnosis of puerperal ovarian vein thrombosis. Obstet Gynecol 1984; 63:61-64

# A Human Case of Monocytic **Ehrlichiosis With Adult Respiratory Distress** Syndrome in Northern California

DUC J. VUGIA, MD, MPH Berkeley, California

ERIC HOLMBERG, MD Point Reyes Station, California

ERIKA M. STEFFE, MD Santa Rosa, California

MICHAEL S. ASCHER, MD DANA GALLO Berkeley, California

IN THE UNITED STATES, human monocytic ehrlichiosis is an emerging tick-borne infectious disease caused by Ehrlichia chaffeensis, a newly identified rickettsial agent that infects mainly mononuclear phagocytes.<sup>13</sup> Recently a human granulocytic ehrlichiosis was documented that is caused by an as-yet-unnamed Ehrlichia species (closely related to Ehrlichia equi) that infects blood granulocytes.35 In both types of human ehrlichiosis, reported cases have often been in older men with a recent history of tick bite or

(Vugia DJ, Holmberg E, Steffe EM, Ascher MS, Gallo D: A human case of monocytic ehrlichiosis with adult respiratory distress syndrome in northern California. West J Med 1996; 164: 525-528)

From the California Department of Health Services, Berkeley, and the Point Reyes Clinic, Point Reyes Station, California. Dr Steffe is in private practice in Santa Rosa, California

Reprint requests to Duc J. Vugia, MD, MPH, Disease Investigations and Surveillance Branch, Division of Communicable Disease Control, California Department of Health Services, 2151 Berkeley Way, Berkeley, CA 94704.

## **ABBREVIATIONS USED IN TEXT**

ARDS = adult respiratory distress syndrome CDC = Centers for Disease Control and Prevention Ig = immunoglobulin

exposure, and the initial illness was usually nonspecific with fever, chills, headache, malaise, myalgia, and nausea.<sup>3,5,8</sup> Laboratory testing often shows abnormalities such as leukopenia, thrombocytopenia, and elevated liver enzyme levels.<sup>3,5,8</sup>

Disease due to *E chaffeensis* was first documented in 1986,9 and to date about 400 cases have been diagnosed from 30 states, with most cases located in the southern United States.367 Asymptomatic *E chaffeensis* infection can occur.10 Disease due to the newer human granulocytic *Ehrlichia* species was only recently documented, with cases from Wisconsin, Minnesota, Connecticut, and other northeastern states.358,11 We report a probable human case of monocytic ehrlichiosis acquired in northern California.

# Report of a Case

The patient, a 64-year-old man, was previously healthy without chronic medical problems. On April 1, 1994, he presented to a rural clinic in Marin County, California, because he had progressive fever, shaking chills, headache, myalgia, nausea, and malaise. On physical examination, his temperature was 39.8°C (103.6°F), and fine bibasilar rales were noted. A borderline interstitial pattern in the basilar areas was noted on a chest x-ray film. He was started on a regimen of oral cephalosporin and sent home pending blood test results.

The following day, when the patient's laboratory tests revealed a platelet count of  $55 \times 10^9$  per liter (55,000 per mm<sup>3</sup>), he was admitted to a hospital. On admission, he had a temperature of 39.2°C (102.6°F), a blood pressure of 140/90 mm of mercury, a pulse rate of 84 beats per minute, and a respiratory rate of 22 per minute, along with persistent rales in both lung bases. There was no cough, dyspnea, jaundice, rash, petechiae, or hepatosplenomegaly. He was alert and coherent but had difficulty with concentration and memory. Admission laboratory studies revealed a platelet count of  $35 \times 10^9$ per liter, a leukocyte count of  $5.7 \times 10^9$  per liter with a differential count of 0.55 (55%) band and 0.31 (31%) segmented forms, and a hemoglobin level of 142 grams per liter (14.2 grams per dl). A total bilirubin level was 47.9 μmol per liter (2.8 mg per dl), compared with 17.1 μmol per liter (1.0 mg per dl) the day before; the serum aspartate aminotransferase level was 87 U per liter, and the lactate dehydrogenase level was 1,171 U per liter. A prothrombin time, activated partial thromboplastin time, and fibrin-split products were within normal limits.

Despite aggressive antimicrobial therapy, his pulmonary function deteriorated progressively. He was transferred to the intensive care unit on the third hospital day and an endotracheal tube introduced on the fourth day. His clinical course and chest film findings were compatible with the adult respiratory distress syndrome (ARDS)

TABLE 1.—Acute, Convalescent, and Follow-up Indirect Immunofluorescence Antibody Titers for Ehrlichia chaffeensis in Case-Patient, Northern California, 1994

Laboratory and	Dates Serum Specimens Obtained		
Antibody Test	4/4	4/19	6/24
California VRDL			
IgG	<1:8	1:32	1:128
CDC			
IgG	<1:16	≥1:512	1:256
IgM	1:32	≥1:512	<1:16
CDC = Centers for Disease Control and Polickettsial Disease Laboratory	revention, Ig = i	mmunoglobulin, V	RDL = Viral and

(Figure 1). Following the placement of a Swan-Ganz catheter and appropriate fluid volume management, his pulmonary function gradually improved. He was extubated on the 22nd day and discharged home on the 26th day of his hospital stay. On discharge, mild residual infiltrates were noted on a chest film and had resolved on follow-up two months later.

Treatment with ceftizoxime sodium and clarithromycin was initially started for "broad-spectrum" coverage, including gram-negative sepsis and bacterial pneumonia. When the patient was transferred to the intensive care unit, his antimicrobial drugs were intravenous ceftizoxime, vancomycin, gentamicin sulfate, and erythromycin, the last one subsequently changed to doxycycline on the fourth hospital day for possible rickettsial agents. Within hours after the regimen of doxycycline was instituted, the patient defervesced and remained afebrile for 12 hours, the first afebrile period since admission. He had fever again the next day, however, and experienced intermittent low-grade fever for the remainder of his hospital stay despite a later addition of metronidazole and fluconazole for possible anaerobic and fungal suprainfection. He did not defervesce entirely until the endotracheal tube was removed, his total parenteral nutrition discontinued, and most of his antimicrobial agents terminated on the 22nd hospital day.

Culture of cerebrospinal fluid and repeated cultures of urine and blood were negative for pathogens, and cultures of aspirate specimens from the endotracheal tube yielded *Candida* species. Endotracheal tube and nasal aspirates were negative for *Legionella pneumophila* and influenza A and B, respectively, by direct fluorescent antibody test. Acute and convalescent sera were negative for immunoglobulin (Ig) G antibodies to *Rickettsia typhi*, *Rickettsia rickettsii*, and *Coxiella burnetii* using standardized procedures. Immunoglobulin G antibodies to *E chaffeensis*, however, showed a fourfold rise (Table 1) by the immunofluorescence method using antigen provided by the Centers for Disease Control and Prevention (CDC). A review of blood smears did not reveal intraleukocytic inclusion bodies.

These sera were also forwarded to the CDC where *E chaffeensis* IgG¹ and IgM (J. Dawson, CDC, Atlanta, Georgia, oral communication, January 1996) results further confirmed an incidence of recent infection (Table 1).

No hantavirus antibodies were detected. Serologic testing for E equi antibodies, the serologic confirmation for human granulocytic ehrlichiosis, at the University of California, Davis, was negative.

Other relevant clinical aspects of the patient's illness included hematologic, hepatic, and neurologic abnormalities. The platelet count declined to  $33 \times 10^9$  per liter, and his leukocyte count reached a nadir of  $2.2 \times 10^9$  per liter on the fourth hospital day. After a course of intravenous doxycycline was initiated, his platelet and leukocyte counts improved and had returned to normal by the sixth hospital day. The aspartate aminotransferase level peaked at 134 U per liter, and his total bilirubin level rose to 63.3 µmol per liter (3.7 mg per dl) before falling. Neurologically, the patient became profoundly disoriented and agitated by the third hospital day. Cerebrospinal fluid glucose and protein concentrations and cell counts were normal, as was computed tomography of his brain. He improved slowly and was fully oriented a month after discharge, but had difficulty remembering details of his hospital stay.

Of epidemiologic importance, the patient frequently hiked three to four miles in the rural coastal area near his weekend home in Marin County. He reported receiving a tick bite on the right side of his neck in mid-March, about two weeks before his illness began, and distinctly remembered the tick bleeding when it was removed and discarded. He had no pets. Five months before the onset of his illness, he traveled to New York City and Washington, DC, but did not remember any tick bite or exposure during that trip.

# Discussion

To our knowledge, this is the first reported human case of ehrlichiosis acquired in California. The patient's serologic test results indicated infection with E chaffeensis or a closely related Ehrlichia species and not the recently discovered human granulocytic ehrlichiosis as antibody to E equi was not demonstrated. He most likely acquired the infection in northern California, probably from the reported tick bite.

His clinical presentation and illness, except for ARDS, was typical of monocytic ehrlichiosis due to E chaffeensis infection described previously in the southern United States. 67 He had nonspecific symptoms initially, and his admission laboratory tests revealed thrombocytopenia and elevated liver enzyme levels; leukopenia developed soon after he was admitted to a hospital. After a regimen of doxycycline was instituted, these abnormalities resolved. Though the patient's fever resolved initially with the use of doxycycline, its intermittent return might have been due to one or a combination of factors other than his ehrlichiosis, such as total parenteral nutrition, atelectasis, or drug allergy. The neurologic involvement was not unusual; confusion has been noted in as much as 29% of patients with ehrlichiosis, clearing slowly with appropriate treatment.<sup>67</sup>

Of clinical interest was his ARDS, which has been noted but not commonly in previous reports of ehrlichiosis. In a recent review of 237 cases of human ehrlichiosis, only 7 of the patients (3%) had pulmonary infiltrates. In

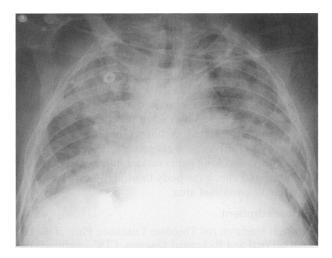


Figure 1.—A chest radiograph shows diffuse bilateral infiltrates consistent with the adult respiratory distress syndrome.

an earlier review of *E chaffeensis* cases, 14 of 32 patients (44%) who had chest roentgenograms taken had evidence of pulmonary infiltrates and 4 of 38 patients (11%) had pulmonary edema; 7 required intubation and mechanical ventilation. In a recent report of cases of human granulocytic ehrlichiosis, two patients who died had pulmonary infiltrates on chest films, one of whom had ARDS.5 Pulmonary hemorrhage and secondary pneumonias due to opportunistic pathogens have been reported in fatal cases, 8,11,13,14 but were not fully evaluated in this case because the patient recovered.

Probable vectors for *E chaffeensis* in the southern United States include the ticks Amblyomma americanum and Dermacentor variabilis,15 whereas those for the human granulocytic ehrlichiosis agent in the northeastern United States include the deer tick, Ixodes scapularis, which is also the tick vector for Lyme disease in that region.16 In California, the tick vector for either agent is unknown, but Amblyomma species are not present. D variabilis ticks are in this state, however, and may be possible vectors for the monocytic ehrlichiosis that occurred in this patient.

This case report adds monocytic ehrlichiosis to the list of tick-borne infectious diseases in California. The clinical presentation is similar to that of Rocky Mountain spotted fever, another rare tick-borne infection in California. Whereas a rash is noted in as much as 90% of patients with Rocky Mountain spotted fever, it is present in less than half of those with monocytic ehrlichiosis (and rarely in patients with human granulocytic ehrlichiosis). Because doxycycline or tetracycline is the antimicrobial drug of choice for both ehrlichiosis and Rocky Mountain spotted fever, its use should be started early in suspected cases while awaiting laboratory confirmation.3 The laboratory diagnosis for both types of Ehrlichia species is more commonly made serologically, although a thorough examination of initial blood smears may reveal intraleukocytic inclusions called morulae, more often with human granulocytic ehrlichiosis than with monocytic ehrlichiosis. 1,3-5,8

Polymerase chain reaction can provide the diagnosis during acute disease, but the test is currently available in only a few research centers. In California, Rocky Mountain spotted fever and monocytic and granulocytic ehrlichioses may be confirmed by sending acute and convalescent sera to the state's Viral and Rickettsial Disease Laboratory.

Ehrlichiosis, Lyme disease, and other tick-borne infections can be prevented by taking precautions to reduce tick exposure. Clinicians should remind their patients to avoid tick-infested areas, wear protective clothing (long-sleeved shirts and long pants tucked into boots), use tick repellents, and check the body thoroughly for ticks after being in a tick-infested area.

## Acknowledgment

Joseph Singleton and Theodore Tzianabos, PhD, of the Division of Viral and Rickettsial Diseases, CDC, confirmed the *Ehrlichia chaffeensis* serology; and John E. Madigan, DVM, of the School of Veterinary Medicine, University of California, Davis, confirmed the *Ehrlichia equi* serology.

#### **REFERENCES**

- 1. Dawson JE, Anderson BE, Fishbein DB, et al: Isolation and characterization of an *Ehrlichia* sp from a patient diagnosed with human ehrlichiosis. J Clin Microbiol 1991; 29:2741-2745
- Anderson BE, Dawson JE, Jones DC, Wilson KH: Ehrlichia chaffeensis, a new species associated with human ehrlichiosis. J Clin Microbiol 1991; 29:2838-2842
- 3. Dumler JS, Bakken JS: Ehrlichial diseases of humans: Emerging tick-borne infections. Clin Infect Dis 1995; 20:1102-1110

- 4. Chen SM, Dumler S, Bakken JS, Walker DH: Identification of a granulocytotropic ehrlichia species as the etiologic agent of human disease. J Clin Microbiol 1994; 32:589-595
- 5. Bakken JS, Dumler JS, Chen SM, Eckman MR, Van Etta LL, Walker DH: Human granulocytic ehrlichiosis in the upper midwest United States: A new species emerging? JAMA 1994; 272:212-218
- 6. Eng TR, Harkness JR, Fishbein DB, et al: Epidemiologic, clinical, and laboratory findings of human ehrlichiosis in the United States, 1988. JAMA 1990; 264:2251-2258
- 7. Fishbein DB, Dawson JE, Robinson LE: Human ehrlichiosis in the United States, 1985-1990. Ann Intern Med 1994; 120:736-743
- 8. Bakken JS, Krueth J, Wilson-Nordskog C, Tilden RL, Asanovich K, Dumler JS: Clinical and laboratory characteristics of human granulocytic ehrlichiosis. JAMA 1996; 275:199-205
- 9. Maeda K, Markowitz N, Hawley RC, Ristic M, Cox D, McDade JE: Human interction with *Ehrlichia canis*, a leukocytic rickettsia. N Engl J Med 1987; 316:853-856
- 10. Yevich SJ, Sanchez JL, DeFraites RF, et al: Seroepidemiology of infections due to spotted fever group rickettsia and *Ehrlichia* in military personnel exposed in areas of the United States where such infections are endemic. J Infect Dis 1995; 171:1266-1273
- 11. Hardalo CJ, Quagliarello V, Dumler JS: Human granulocytic ehrlichiosis in Connecticut: Report of a fatal case. Clin Infect Dis 1995; 21:910-914
- 12. McDade JE, Fishbein DB: Rickettsiaceae—The rickettsiae, *In Balows A*, Hausler WJ Jr, Lennette EH (Eds): Laboratory Diagnosis of Infectious Diseases—Principles and Practices. New York, NY, Springer-Verlag, 1988, pp 864-890
- 13. Paddock CD, Suchard DP, Grumbach KL, et al: Brief report: Fatal seronegative ehrlichiosis in a patient with HIV infection. N Engl J Med 1993; 329:1164-1167
- Dumler JS, Sutker WL, Walker DH: Persistent infection with Ehrlichia chaffeensis. Clin Infect Dis 1993; 17:903-905
- 15. Anderson BE, Sims KG, Olson JG, et al: *Amblyomma americanum*: A potential vector of human ehrlichiosis. Am J Trop Med Hyg 1993; 49:239-244
- 16. Magnarelli LA, Stafford KC III, Mather TN, Yeh M, Horn KD, Dumler JS: Hemocytic rickettsia-like organisms in ticks: Serologic reactivity with antisera to ehrlichiae and detection of DNA of agent of human granulocytic ehrlichiosis by PCR. J Clin Microbiol 1995; 33:2710-2714